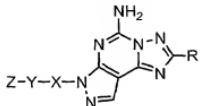


We claim:

1. Compounds having the structural formula

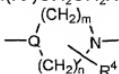


5 or a pharmaceutically acceptable salt thereof, wherein

R is R<sup>1</sup>-furanyl, R<sup>1</sup>-thienyl, R<sup>1</sup>-pyridyl, R<sup>1</sup>-pyridyl N-oxide, R<sup>1</sup>-oxazolyl, R<sup>10</sup>-phenyl, R<sup>1</sup>-pyrrolyl or C<sub>4</sub>-C<sub>6</sub> cycloalkenyl;

X is C<sub>2</sub>-C<sub>6</sub> alkylene or -C(O)CH<sub>2</sub>-;

Y is -N(R<sup>2</sup>)CH<sub>2</sub>CH<sub>2</sub>N(R<sup>3</sup>)-, -OCH<sub>2</sub>CH<sub>2</sub>N(R<sup>2</sup>)-, -O-, -S-, -CH<sub>2</sub>S-, -(CH<sub>2</sub>)<sub>2</sub>-NH-, or



and

Z is R<sup>5</sup>-phenyl, R<sup>5</sup>-phenyl(C<sub>1</sub>-C<sub>6</sub>)alkyl, R<sup>5</sup>-heteroaryl, diphenylmethyl, R<sup>6</sup>-C(O)-,



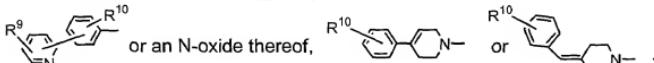
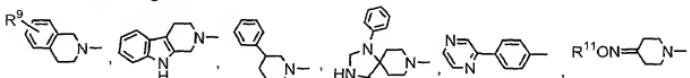
R<sup>5</sup>-SO<sub>2</sub>-, R<sup>6</sup>-OC(O)-, R<sup>7</sup>-N(R<sup>8</sup>)-C(O)-, R<sup>7</sup>-N(R<sup>8</sup>)-C(S)-, phenyl-CH(OH)-, or



phenyl-C(=NOR<sup>2</sup>)-; or when Q is H, Z is also phenylamino or pyridylamino;

15 or

Z and Y together are



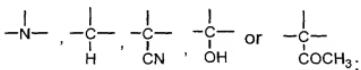
R<sup>1</sup> is 1 to 3 substituents independently selected from hydrogen, C<sub>1</sub>-C<sub>6</sub>-alkyl,

20 -CF<sub>3</sub>, halogen, -NO<sub>2</sub>, -NR<sup>12</sup>R<sup>13</sup>, C<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkylsulfinyl, and C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl;

R<sup>2</sup> and R<sup>3</sup> are independently selected from the group consisting of hydrogen and C<sub>1</sub>-C<sub>6</sub> alkyl;

m and n are independently 2-3;

Q is



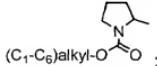
R<sup>4</sup> is 1-2 substituents independently selected from the group consisting of hydrogen and C<sub>1</sub>-C<sub>6</sub>alkyl, or two R<sup>4</sup> substituents on the same carbon can form =O;

R<sup>5</sup> is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C<sub>1</sub>-C<sub>6</sub> alkyl, hydroxy, C<sub>1</sub>-C<sub>6</sub> alkoxy, -CN, di-((C<sub>1</sub>-C<sub>6</sub>)alkyl)amino, -CF<sub>3</sub>, -OCF<sub>3</sub>, acetyl, -NO<sub>2</sub>, hydroxy(C<sub>1</sub>-C<sub>6</sub>)alkoxy, (C<sub>1</sub>-C<sub>6</sub>)-alkoxy(C<sub>1</sub>-C<sub>6</sub>)alkoxy, di-((C<sub>1</sub>-C<sub>6</sub>)-alkoxy)(C<sub>1</sub>-C<sub>6</sub>)alkoxy, (C<sub>1</sub>-C<sub>6</sub>)-alkoxy(C<sub>1</sub>-C<sub>6</sub>)alkoxy-(C<sub>1</sub>-C<sub>6</sub>)-alkoxy, carboxy(C<sub>1</sub>-C<sub>6</sub>)-alkoxy, (C<sub>1</sub>-C<sub>6</sub>)-alkoxycarbonyl(C<sub>1</sub>-C<sub>6</sub>)alkoxy, (C<sub>3</sub>-C<sub>6</sub>)cycloalkyl(C<sub>1</sub>-C<sub>6</sub>)alkoxy,

10      di-((C<sub>1</sub>-C<sub>6</sub>)alkyl)amino(C<sub>1</sub>-C<sub>6</sub>)alkoxy, morpholinyl, (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub><sup>-</sup>, (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sup>-</sup>(C<sub>1</sub>-C<sub>6</sub>)alkoxy, tetrahydropyranloxy, (C<sub>1</sub>-C<sub>6</sub>)alkylcarbonyl(C<sub>1</sub>-C<sub>6</sub>)-alkoxy, (C<sub>1</sub>-C<sub>6</sub>)-alkoxycarbonyl, (C<sub>1</sub>-C<sub>6</sub>)alkylcarbonyloxy(C<sub>1</sub>-C<sub>6</sub>)-alkoxy, -SO<sub>2</sub>NH<sub>2</sub>, phenoxy,

15       $\begin{array}{c} (\text{C}_1\text{-}\text{C}_6 \text{ alkyl}) \\ | \\ -C=NR^2 \end{array}$ , ; or adjacent R<sup>5</sup> substituents together are -O-CH<sub>2</sub>-O-, -O-CH<sub>2</sub>CH<sub>2</sub>-O-, -O-CF<sub>2</sub>-O- or -O-CF<sub>2</sub>CF<sub>2</sub>-O- and form a ring with the carbon atoms to which they are attached;

20      R<sup>6</sup> is (C<sub>1</sub>-C<sub>6</sub>)alkyl, R<sup>6</sup>-phenyl, R<sup>6</sup>-phenyl(C<sub>1</sub>-C<sub>6</sub>)alkyl, thienyl, pyridyl, (C<sub>3</sub>-C<sub>6</sub>)-cycloalkyl, (C<sub>1</sub>-C<sub>6</sub>)alkyl-OC(O)-NH-(C<sub>1</sub>-C<sub>6</sub>)alkyl-, di-((C<sub>1</sub>-C<sub>6</sub>)alkyl)aminomethyl, or



R<sup>7</sup> is (C<sub>1</sub>-C<sub>6</sub>)alkyl, R<sup>6</sup>-phenyl or R<sup>6</sup>-phenyl(C<sub>1</sub>-C<sub>6</sub>)alkyl;

25      R<sup>8</sup> is hydrogen or C<sub>1</sub>-C<sub>6</sub> alkyl; or R<sup>7</sup> and R<sup>8</sup> together are -(CH<sub>2</sub>)<sub>p</sub>-A-(CH<sub>2</sub>)<sub>q</sub>, wherein p and q are independently 2 or 3 and A is a bond, -CH<sub>2</sub>-, -S- or -O-, and form a ring with the nitrogen to which they are attached;

R<sup>9</sup> is 1-2 groups independently selected from hydrogen, C<sub>1</sub>-C<sub>6</sub> alkyl, hydroxy, C<sub>1</sub>-C<sub>6</sub> alkoxy, halogen, -CF<sub>3</sub> and (C<sub>1</sub>-C<sub>6</sub>)alkoxy(C<sub>1</sub>-C<sub>6</sub>)alkoxy ;

30      R<sup>10</sup> is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C<sub>1</sub>-C<sub>6</sub> alkyl, hydroxy, C<sub>1</sub>-C<sub>6</sub> alkoxy, -CN, -NH<sub>2</sub>, C<sub>1</sub>-C<sub>6</sub>alkylamino, di-((C<sub>1</sub>-C<sub>6</sub>)alkyl)amino, -CF<sub>3</sub>, -OCF<sub>3</sub> and -S(O)<sub>2</sub>(C<sub>1</sub>-C<sub>6</sub>)alkyl;

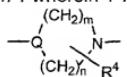
R<sup>11</sup> is H, C<sub>1</sub>-C<sub>6</sub> alkyl, phenyl, benzyl, C<sub>2</sub>-C<sub>6</sub> alkenyl, C<sub>1</sub>-C<sub>6</sub> alkoxy(C<sub>1</sub>-C<sub>6</sub>)alkyl, di-((C<sub>1</sub>-C<sub>6</sub>)alkyl)amino(C<sub>1</sub>-C<sub>6</sub>)alkyl, pyrrolidinyl(C<sub>1</sub>-C<sub>6</sub>)alkyl or piperidino(C<sub>1</sub>-C<sub>6</sub>)alkyl;

R<sup>12</sup> is H or C<sub>1</sub>-C<sub>6</sub> alkyl; and  
R<sup>13</sup> is (C<sub>1</sub>-C<sub>6</sub>)alkyl-C(O)- or (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub><sup>-</sup>.

2. A compound of claim 1 wherein R is R<sup>1</sup>-furanyl.

- 5 3. A compound of claim 1 wherein X is C<sub>2</sub>-C<sub>6</sub> alkylene.

4. A compound of claim 1 wherein Y is



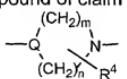
- 10 5. A compound of claim 5 wherein Q is  $\begin{matrix} | \\ -N- \end{matrix}$  or  $\begin{matrix} | \\ -CH- \end{matrix}$ .

6. A compound of claim 5 wherein m and n are each 2, and R<sup>4</sup> is H.

- 15 7. A compound of claim 1 wherein Z is R<sup>5</sup>-phenyl, R<sup>5</sup>-heteroaryl, R<sup>6</sup>-C(O)- or R<sup>6</sup>-SO<sub>2</sub><sup>-</sup>.

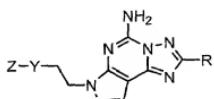
8. A compound of claim 7 wherein R<sup>5</sup> is H, halogen, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkoxy, hydroxy(C<sub>1</sub>-C<sub>6</sub>)alkoxy or (C<sub>1</sub>-C<sub>6</sub>)alkoxy(C<sub>1</sub>-C<sub>6</sub>)alkoxy, or R<sup>6</sup> is R<sup>5</sup>-phenyl.

- 20 9. A compound of claim 1 wherein R is R<sup>1</sup>-furanyl, X is C<sub>2</sub>-C<sub>6</sub> alkylene, Y is



- Q is  $\begin{matrix} | \\ -N- \end{matrix}$  or  $\begin{matrix} | \\ -CH- \end{matrix}$ , m and n are each 2, R<sup>4</sup> is H, Z is R<sup>5</sup>-phenyl, R<sup>5</sup>-heteroaryl, R<sup>6</sup>-C(O)- or R<sup>6</sup>-SO<sub>2</sub><sup>-</sup>, R<sup>5</sup> is H, halogen, C<sub>1</sub>-C<sub>6</sub> alkyl, C<sub>1</sub>-C<sub>6</sub> alkoxy, hydroxy(C<sub>1</sub>-C<sub>6</sub>)alkoxy or (C<sub>1</sub>-C<sub>6</sub>)alkoxy(C<sub>1</sub>-C<sub>6</sub>)alkoxy, and R<sup>6</sup> is R<sup>5</sup>-phenyl.

- 25 10. A compound of claim 1 selected from the group consisting of compounds of the formula



- 30 30 wherein R and Z-Y are as defined in the following table:

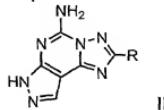
Z-Y-	R
<chem>Fc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>
<chem>c1ccncc1N2CCNCC2</chem>	<chem>C=C1OC1</chem>
<chem>Fc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>
<chem>CCOCc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>
<chem>CCOCc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>
<chem>CC(=O)Oc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>
<chem>CC(=O)Oc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>
<chem>Clc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>
<chem>CCOCc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>
<chem>Fc1cnc2c(N3CCNCC3)nc2c1</chem>	<chem>C=C1OC1</chem>
<chem>CH3Cc1cnc2c(N3CCNCC3)nc2c1</chem>	<chem>C=C1OC1</chem>
<chem>Fc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>
<chem>Fc1ccc(F)c(N2CCNCC2)c1</chem>	<chem>C=C1OC1</chem>

11. A pharmaceutical composition comprising a therapeutically effective amount of a compound of claim 1 in a pharmaceutically acceptable carrier.
- 5 12. A method of treating central nervous system diseases or stroke, comprising administering an effective amount of a compound of formula I to a mammal in need of such treatment.

13. A method of claim 12 for treating depression, cognitive diseases and neurodegenerative diseases.

5 14. A method of claim 13 for treating Parkinson's disease, senile dementia or psychoses of organic origin.

15. A process of preparing a compound of formula II



wherein R is R<sup>1</sup>-furanyl, R<sup>1</sup>-thienyl, R<sup>1</sup>-pyridyl, R<sup>1</sup>-pyridyl N-oxide, R<sup>1</sup>-oxazolyl, 10 R<sup>10</sup>-phenyl, R<sup>1</sup>-pyrrolyl or C<sub>4</sub>-C<sub>6</sub> cycloalkenyl;

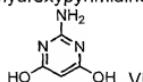
R<sup>1</sup> is 1 to 3 substituents independently selected from hydrogen, C<sub>1</sub>-C<sub>6</sub>-alkyl, -CF<sub>3</sub>, halogen, -NO<sub>2</sub>, -NR<sup>12</sup>R<sup>13</sup>, C<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkylsulfinyl, and C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl;

15 R<sup>10</sup> is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C<sub>1</sub>-C<sub>6</sub> alkyl, hydroxy, C<sub>1</sub>-C<sub>6</sub> alkoxy, -CN, -NH<sub>2</sub>, C<sub>1</sub>-C<sub>6</sub> alkylamino, di-((C<sub>1</sub>-C<sub>6</sub>)alkyl)amino, -CF<sub>3</sub>, -OCF<sub>3</sub> and -S(O)<sub>0-2</sub>(C<sub>1</sub>-C<sub>6</sub>)alkyl;

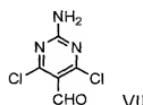
R<sup>12</sup> is H or C<sub>1</sub>-C<sub>6</sub> alkyl; and

15 R<sup>13</sup> is (C<sub>1</sub>-C<sub>6</sub>)alkyl-C(O)- or (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub>-;  
comprising

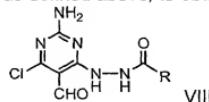
20 (1) treating 2-amino-4,6-dihydroxypyrimidine



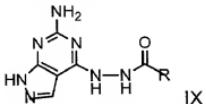
with POCl<sub>3</sub> in dimethylformamide to obtain 2-amino-4,6-dichloropyrimidine-5-carboxaldehyde



25 (2) treating carboxaldehyde VII with a hydrazide of the formula H<sub>2</sub>N-NH-C(O)-R, wherein R is as defined above, to obtain

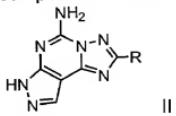


(3) treating the intermediate of formula VIII with hydrazine hydrate to form a pyrazolo ring, thus obtaining the intermediate of formula IX



5 (4) forming the desired compound of formula II by dehydrative rearrangement.

16. A process for preparing a compound of the formula II



wherein R is R<sup>1</sup>-furanyl, R<sup>1</sup>-thienyl, R<sup>1</sup>-pyridyl, R<sup>1</sup>-pyridyl N-oxide, R<sup>1</sup>-oxazolyl, R<sup>10</sup>-phenyl, R<sup>1</sup>-pyrrolyl or C<sub>4</sub>-C<sub>6</sub> cycloalkenyl;

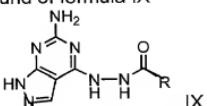
10 R<sup>1</sup> is 1 to 3 substituents independently selected from hydrogen, C<sub>1</sub>-C<sub>6</sub>-alkyl, -CF<sub>3</sub>, halogen, -NO<sub>2</sub>, -NR<sup>12</sup>R<sup>13</sup>, C<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkylsulfinyl, and C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl;

15 R<sup>10</sup> is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C<sub>1</sub>-C<sub>6</sub> alkyl, hydroxy, C<sub>1</sub>-C<sub>6</sub> alkoxy, -CN, -NH<sub>2</sub>, C<sub>1</sub>-C<sub>6</sub> alkylamino, di-((C<sub>1</sub>-C<sub>6</sub>)alkyl)amino, -CF<sub>3</sub>, -OCF<sub>3</sub> and -S(O)<sub>0-2</sub>(C<sub>1</sub>-C<sub>6</sub>)alkyl;

R<sup>12</sup> is H or C<sub>1</sub>-C<sub>6</sub> alkyl; and

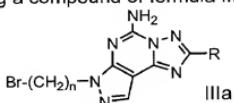
R<sup>13</sup> is (C<sub>1</sub>-C<sub>6</sub>)alkyl-C(O)- or (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub>-;

comprising converting a compound of formula IX



20 into the desired compound of formula II by dehydrative rearrangement.

17. A process for preparing a compound of formula IIIa



wherein R is R<sup>1</sup>-furanyl, R<sup>1</sup>-thienyl, R<sup>1</sup>-pyridyl, R<sup>1</sup>-pyridyl N-oxide, R<sup>1</sup>-oxazolyl, R<sup>10</sup>-phenyl, R<sup>1</sup>-pyrrolyl or C<sub>4</sub>-C<sub>6</sub> cycloalkenyl;

25 R<sup>1</sup> is 1 to 3 substituents independently selected from hydrogen, C<sub>1</sub>-C<sub>6</sub>-alkyl, -CF<sub>3</sub>, halogen, -NO<sub>2</sub>, -NR<sup>12</sup>R<sup>13</sup>, C<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>6</sub> alkylthio, C<sub>1</sub>-C<sub>6</sub> alkylsulfinyl, and C<sub>1</sub>-C<sub>6</sub> alkylsulfonyl;

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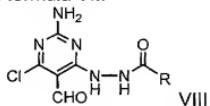
R<sup>10</sup> is 1 to 5 substituents independently selected from the group consisting of hydrogen, halogen, C<sub>1</sub>-C<sub>6</sub> alkyl, hydroxy, C<sub>1</sub>-C<sub>6</sub> alkoxy, -CN, -NH<sub>2</sub>, C<sub>1</sub>-C<sub>6</sub>alkylamino, di-((C<sub>1</sub>-C<sub>6</sub>)alkyl)amino, -CF<sub>3</sub>, -OCF<sub>3</sub> and -S(O)<sub>0-2</sub>(C<sub>1</sub>-C<sub>6</sub>)alkyl;

R<sup>12</sup> is H or C<sub>1</sub>-C<sub>6</sub> alkyl; and

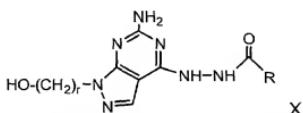
5 R<sup>13</sup> is (C<sub>1</sub>-C<sub>6</sub>)alkyl-C(O)- or (C<sub>1</sub>-C<sub>6</sub>)alkyl-SO<sub>2</sub>-;

comprising

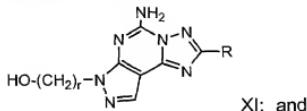
(1) treating a chloride of formula VIII



with a hydroxyalkyl hydrazine of the formula HO-(CH<sub>2</sub>)<sub>r</sub>-NHNH<sub>2</sub>, wherein r is 2-6, to obtain



(2) cyclizing the intermediate of formula X by dehydrative rearrangement to obtain the tricyclic intermediate of formula XI



15 (3) converting the hydroxy compound of formula XI to the bromide of formula IIIa.

18. A pharmaceutical composition comprising a therapeutically effective amount of a combination of a compound of claim 1 and 1 to 3 other agents useful in treating

20 Parkinson's disease in a pharmaceutically acceptable carrier

19. A method of treating Parkinson's disease comprising administering to a mammal in need of such treatment an effective amount of a combination of a compound of claim 1 and 1 to 3 other agents useful in treating Parkinson's disease.

25

20. The method of claim 19 wherein the other agents are selected from the group consisting of L-DOPA, dopaminergic agonists, MAO-B inhibitors, DOPA decarboxylase inhibitors and COMT inhibitors.

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